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For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.

A3

(54) Title: MUCUNA PRURIENS AND EXTRACTS THEREOF FOR THE TREATMENT OF NEUROLOGICAL DISEASES

(57) Abstract: The present invention provides pharmaceutical compositions comprising Mucuna pruriens seeds or one or more Mucuna pruriens seed components, substances, fractions or mixtures or substances obtained therefrom. Furthermore, the invention relates to the use of Mucuna pruriens seed powder or one or more Mucuna pruriens components, substances, fractions or mixtures or substances obtained therefrom for the preparation of a pharmaceutical composition for preventing, alleviating or treating neurological diseases. Additionally, the invention relates to the use of Mucuna pruriens seeds for the preparation of a pharmaceutical composition for neuroprotection or neurostimulation and to methods of preparing extracts of Mucuna pruriens which can be used for the preparation of a pharmaceutical composition for treating neurological diseases. Finally, the invention relates to the use of Mucuna pruriens seeds for the preparation of a pharmaceutical composition for the treatment of Parkinson's Disease to obtain a broader therapeutic window in L-Dopa therapy, to delay a need for combination therapy, to obtain an earlier onset and longer duration of L-Dopa efficacy, and to prevent or alleviate acute and chronic L-Dopa toxicity.



AL SEARCH REPORT

A. CLASSIFICATION OF SUBJECT MATTER IPC 7 A61K35/78 A61P25/16

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols) $\begin{tabular}{ll} IPC & 7 & A61K & A61P \end{tabular}$

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

EPO-Internal, WPI Data, EMBASE, MEDLINE, BIOSIS, PAJ, COMPENDEX

C DOCUM	ENTS CONSIDERED TO BE RELEVANT		
Category °	Citation of document, with indication, where appropriate, of the	relevant passages	Relevant to claim No.
X	US 6 106 839 A (PRUTHY PUNEET 22 August 2000 (2000-08-22) abstract	1-20, 26-30	
X	US 3 666 802 A (ADAMS PHILLIP E 30 May 1972 (1972-05-30) abstract column 1, line 6 - line 10	1-20, 26-30	
X	VAIDYA A B ET AL: "Treatment o Parkinson's disease with the co - Mucuna pruriens Bak" NEUROLOGY INDIA 1978 INDIA, vol. 26, no. 4, 1978, pages 171 XP009024725 Introduction & Discussion	whage plant	1-20, 26-30
X Furti	ner documents are listed in the continuation of box C.	χ Patent family members are listed in	n annex.
"A" docume consid "E" earlier of filing d "L" docume which citatior "O" docume other r "P" docume later th	ant which may throw doubts on priority claim(s) or is cited to establish the publication date of another in or other special reason (as specified) and referring to an oral disclosure, use, exhibition or means are published prior to the international filing date but can the priority date claimed	"T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art. "&" document member of the same patent family	
	actual completion of the International search 8 January 2004	Date of malling of the international sear	rch report
	nalling address of the ISA European Patent Office, P.B. 5818 Patentlaan 2 NL – 2280 HV Rijswijk Tel. (+31–70) 340–2040, Tx. 31 651 epo nl, Fax: (+31–70) 340–3016	Authorized officer Pilling, S	



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	ation) DOCUMENTS CONSIDERED TO BE RELEVANT		
Сагедогу Т	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.	
X	HUSSAIN G ET AL: "Mucuna pruriens proves more effective than L-DOPA in Parkinson's disease animal model" PHYTOTHERAPY RESEARCH 1997 UNITED KINGDOM, vol. 11, no. 6, 1997, pages 419-423, XP009024754 ISSN: 0951-418X abstract	1-20, 26-30	
X	"An alternative medicine treatment for Parkinson's disease: results of a multicenter clinical trial. HP-200 in Parkinson's Disease Study Group." JOURNAL OF ALTERNATIVE AND COMPLEMENTARY MEDICINE (NEW YORK, N.Y.) UNITED STATES 1995 FALL, vol. 1, no. 3, October 1995 (1995-10), pages 249-255, XP009024734 ISSN: 1075-5535 abstract	1-20, 26-30	
X	MANYAM BALA V ET AL: "An open efficacy and tolerability study of HP-200 in Parkinson's disease" NEUROLOGY, vol. 44, no. 4 SUPPL. 2, 1994, page A369, XP009024722 46th Annual Meeting of the American Academy of Neurology; Washington, D.C., USA; May 1-7, 1994 ISSN: 0028-3878 abstract	1-20, 26-30	
X	MAHAJANI S S ET AL: "Bioavailability of L-DOPA from HP-200 - A formulation of seed powder of Mucuna pruriens (Bak): A pharmacokinetic and pharmacodynamic study" PHYTOTHERAPY RESEARCH 1996 UNITED KINGDOM, vol. 10, no. 3, 1996, pages 254-256, XP009024724 ISSN: 0951-418X Introduction & Discussion	1-20, 26-30	
X	NAGASHAYANA N ET AL: "ASSOCIATION OF L-DOPA WITH RECOVERY FOLLOWING AYURVEDA MEDICATION IN PARKINSON'S DISEASE" JOURNAL OF NEUROLOGICAL SCIENCES, ELSEVIER SCIENTIFIC PUBLISHING CO, AMSTERDAM, NL, vol. 176, no. 2,	1-20, 26-30	
	15 June 2000 (2000-06-15), pages 124-127, XP001147413 ISSN: 0022-510X abstract		
	-/		



PCT/EP 03/10975

		101/EF 03/109/5
	nation) DOCUMENTS CONSIDERED TO BE RELEVANT	
Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	MANYAM B V ET AL: "Traditional and complementary therapies in Parkinson's disease." ADVANCES IN NEUROLOGY. UNITED STATES 1999, vol. 80, 1999, pages 565-574, XP009024735 ISSN: 0091-3952 page 569, column 2, line 1 - page 570, column 3, line 28	1-20, 26-30

INTERNATIONAL SEARCH REPORT

rational application No. PCT/EP 03/10975

Observations where certain claims were found unsearchable (Continuation of Item 1 of first sheet)				
This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:				
Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:				
2. X Claims Nos.: 1-20, 26 to 30 (part) because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically: see FURTHER INFORMATION sheet PCT/ISA/210				
3. Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).				
Box II Observations where unity of invention is lacking (Continuation of item 2 of first sheet)				
This International Searching Authority found multiple inventions in this international application, as follows:				
see additional sheet				
As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.				
As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.				
As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:				
4. No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.: 1-20, 30 + 26-29 in part				
Remark on Protest The additional search fees were accompanied by the applicant's protest. No protest accompanied the payment of additional search fees.				

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

Continuation of Box I.2

Claims Nos.: 1-20, 26 to 30 (part)

Present Claim 1 relates to a pharmaceutical composition comprising one or more Mucuna pruriens seed COMPONENTS, SUBSTANCES, FRACTIONS or MIXTURES of substances obtained therefrom and a pharmaceutically acceptable diluent, excipient or carrier. Thus Claim 1 appears to define not only a pharmaceutical composition comprising components extracted from Mucuna pruriens but also pharmaceutical compositions comprising ANY SUBSTANCE OBTAINABLE from Mucuna pruriens, e.g. commonly used pharmaceutical substances such as 5-hydroxytryptamine, alanine, arginine or L-DOPA (see present page 4 lines 31 to page 5 line 12). Thus, present Claim 1 appears to define, at least in part, conventional pharmaceutical compositions for which thousands of novelty destroying disclosures may be found. Thus, a search covering the full scope of present Claim 1 is not feasible.

Even a preliminary more restricted search covering pharmaceutical compositions comprising only extracts of Mucuna pruriens revealed many hundreds of disclosures of such pharmaceutical compositions. This appears to confirm the many known traditional uses of extracts of Mucuna pruriens as disclosed by the Applicant in the present description (see present page 5 lines 12 to 17).

Thus, the search was restricted to subject matter for which a search was feasible and for which there appears to be some support in the description, namely the use of Mucuna pruriens for the preparation of pharmaceutical compositions for the treatment of neurological disorders, i.e. Huntington's disease, Alzheimer's disease or Parkinson's disease. The Applicant is warned however, that, in view of the many relevant documents found, that further searching may be necessary at a later stage.

The applicant's attention is drawn to the fact that claims relating to inventions in respect of which no international search report has been established need not be the subject of an international preliminary examination (Rule 66.1(e) PCT). The applicant is advised that the EPO policy when acting as an International Preliminary Examining Authority is normally not to carry out a preliminary examination on matter which has not been searched. This is the case irrespective of whether or not the claims are amended following receipt of the search report or during any Chapter II procedure. If the application proceeds into the regional phase before the EPO, the applicant is reminded that a search may be carried out during examination before the EPO (see EPO Guideline C-VI, 8.5), should the problems which led to the Article 17(2) declaration be overcome.



Intra-nal Application No
PCT/EP 03/10975

Patent do cited in sea		Publication date	Patent fa membe	Publication date
US 6106	839 A	22-08-2000	NONE	
US 3666	802 A	30-05-1972	NONE	